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• Any questions about L.A. Care Health Plan's Provider Continuing Education (PCE) Program and our CME/CE activities, please email Leilanie Mercurio at <u>Imercurio@lacare.org</u>

Presenter's Bio

Fola May, MD, PhD, MPhil, is an Associate Professor of Medicine at the David Geffen School of Medicine at University of California Los Angeles (UCLA), Director of Quality Improvement in the Vatche and Tamar Manoukian



Division of Digestive Diseases, Director of the May Laboratory, Associate Director of the UCLA Kaiser Permanente Center for Health Equity, and staff physician in the Veterans Affairs (VA) Greater Los Angeles Healthcare System.

Dr. May's research is funded by several agencies, including the National Institutes of Health (NIH), and aims to improve population preventive health strategies, increase access to cancer preventive services, and eliminate health disparities domestically and internationally.

An Update on Colorectal Cancer

Folasade P. May M.D., Ph.D., M.Phil. Associate Professor of Medicine University of California Los Angeles Veterans Health Administration September 28, 2023 | WebEx, 12:00 pm – 1:30 pm PST, 1.50 CME / CE Credits Directly Provided CME / CE Activity by L.A. Care Health Plan



Disclosures

Folasade P. May MD PhD MPhil

Consultant/Advisor for:	Kimberly-Clarke/Cottonelle; Takeda; Seed Global Health; Medtronic; Johnson & Johnson; Freenome; Owl Peek Laboratories; Saint Supply
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- Leilanie Mercurio, L.A. Care Provider Continuing Education (PCE) Program Manager, CME Planner
- Bridget Freeley, American Cancer Society Associate Director, State Partnerships, CME Planner

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Presentation Overview

- Colorectal cancer (CRC) is common, even in young adults.
- Screening for CRC is effective.
- Overview of screening guidelines and tests.
- Screening utilization and barriers to screening.
- Achieving equity in CRC.
- Future priorities.





Learning Objectives

At the completion of the activity, learners can:

- 1. Summarize updated colorectal cancer screening guidelines.
- 2. Identify at least 3 symptoms of colorectal cancer.
- 3. Specify recommendations about colorectal cancer screening modalities.
- 4. List at least 3 effective interventions to reduce disparities in colorectal cancer screening.



Colorectal Cancer is Common

<u>#3 cause of cancer in men and women in the United</u>

			Males
Prostate	288,300	29%	
Lung & bronchus	117,550	12%	
Colon & rectum	81,860	8%	
Urinary bladder	62,420	6%	
Melanoma of the skin	58,120	6%	
Kidney & renal pelvis	52,360	5%	
Non-Hodgkin lymphoma	44,880	4%	
Oral cavity & pharynx	39,290	4%	
Leukemia	35,670	4%	
Pancreas	33,130	3%	
All Sites	1,010,310	100%	

Females

All Sites	948,000	100%
Leukemia	23,940	3%
Kidney & renal pelvis	29,440	3%
Pancreas	30,920	3%
Thyroid	31,180	3%
Non-Hodgkin lymphoma	35,670	4%
Melanoma of the skin	39,490	4%
Uterine corpus	66,200	7%
Colon & rectum	71,160	8%
Lung & bronchus	120,790	13%
Breast	297,790	31%



American Cancer Society, Inc., 2023.

Colorectal Cancer is Deadly

<u>#2 cause</u> of cancer-related deaths in the United

			Males
Lung & bronchus	67,160	21%	
Prostate	34,700	11%	
Colon & rectum	28,470	9%	
Pancreas	26,620	8%	
Liver & intrahepatic bile duct	19,000	6%	
Leukemia	13,900	4%	
Esophagus	12,920	4%	
Urinary bladder	12,160	4%	
Non-Hodgkin lymphoma	11,780	4%	
Brain & other nervous system	11,020	3%	
All Sites	322,080	100%	

Females

	Lung & bronchus	59,910	21%
Ļ	Breast	43,170	15%
	Colon & rectum	24,080	8%
	Pancreas	23,930	8%
	Ovary	13,270	5%
	Uterine corpus	13,030	5%
	Liver & intrahepatic bile duct	10,380	4%
	Leukemia	9,810	3%
	Non-Hodgkin lymphoma	8,400	3%
	Brain & other nervous system	7,970	3%
	All Sites	287,740	100%



American Cancer Society, Inc., 2023.

Cases are Rising in Young Adults



There has been a 51% increase in CRC incidence in individuals aged 20-49 since the early 1990s.



SEER 9 database. Delay-adjusted rates, 1975-2012; 2-year averages. Photo courtesy of Rebecca Siegel. Rahib et al., JAMA open network, 2021.

What's the Colon for Anyway?

The colon and rectum:

- Large intestine or large bowel
- · Last parts of the digestive system

The **colon** functions to:

- Absorb water and salt from food Ascending colon
- Form stool

The **rectum** functions to:

Store stool until ready to pass





Everyone is at Risk

Risk Factors You <u>Can Not</u> Change

- Age
- Male genotype
- Race (Black Americans)
- Personal history of polyps
- Family history of polyps or cancer
- Inflammatory bowel disease (IBD)
- Inherited polyp syndromes

Risk Factors You <u>Can</u> Change

• Diet:

- Low fiber diet
- High in animal fat
- Physical inactivity
- Obesity
- Type 2 diabetes
- Tobacco
- Heavy alcohol





Colorectal Cancer Originates as Polyps



years & years & years





Colorectal Cancer



Normal Colon

How can I prevent colorectal cancer?





USPSTF Recommended Screening Modalities (Average-risk individuals)



Direct-visualization techniques





Davidson K, et al. JAMA. 2021;325(19):1965-1977.

Colonoscopy





- **Performed** in a hospital or medical clinic
- Requires bowel preparation
- Gastroenterologist uses a "colonoscope," a long flexible tube with a light at the end
- Requires conscious **sedation** or monitored anesthesia care
- Examines the walls of the colon (20-30 minutes)
- Risks are very small (1:1000) and include bleeding, infection, and colon injury
- Considered the **gold standard** for finding colon cancer or precancerous polyps
- If normal, repeated every 10 years



What is a Colonoscopy Like?

A survey of almost **50,000** patients:

- Most (about 8 out of 10) said the colonoscopy was less uncomfortable than they expected.
- People often agree that "It's not as bad as I thought."



Several studies have found high levels of patient satisfaction and willingness to return under the same conditions.





Ghanouni et al., 2016 Charter et al., 2009.

Fecal Immunochemical Test (FIT)



- Second most common screening test
- Stool-based test that can be performed at home
- Tests the stool for small amounts of blood which *may be* a sign of colon cancer
- Very low-risk screening option
- Must be completed yearly to be effective
- If abnormal, a colonoscopy <u>must</u> be performed to find the source of blood loss



Multitarget Stool DNA (Cologuard)

- Tests stool for 11 pre-cancer and cancer biomarkers and for human hemoglobin (i.e. DNA+FIT)
- 3-year screening interval
- Test failures and false-positive rate 13%
 <u>7 DNA mutation markers</u>
 2 DNA methylation markers









CRC Screening is Effective

Test	Evidence	Certainty of Evidence
Flexible sigmoidoscopy (v. no screening)	Incidence reduction (0.78; 0.74- 0.83) Mortality reduction (0.74; 0.68-0.80)	High
Annual gFOBT (v. no screening)	Incidence reduction (0.81; 0.71- 0.93) Mortality reduction (0.68; 0.56-0.82)	Moderate to High

Guaiac-based FOBT and flexible sigmoidoscopy shown to reduce incidence and mortality in randomized trials



CRC Screening is Effective

Most common applied tests today are the fecal immunochemical test (FIT) and colonoscopy:

- Supported largely by observational data
- Inferred benefit due to gFOBT and flexible sigmoidoscopy data.



Colonoscopy Prevalence (2000 to 2018) and

Survey year/Year of diagnosis



Helsingen, et al. NEJM Evid. 2022, 1. Bretthauer et al, N Engl J Med 2022; 387:1547-1556. Siegel et al, CA Cancer J Clin. 2020 May;70(3):145-164.

CRC Screening is Effective

NORDICC Trial (Poland, Norway, Sweden, Netherlands; 2009-2014)

- 84,000 men and women age 55 to 64.
- PPA: 18% incidence reduction; no mortality reduction.
- ITT: 31% incidence reduction; 50% mortality reduction.
- But ...
 - Only 42% completed colonoscopy
 - Relatively few polyps detected
 - Only 10 years of follow-up
 - US population is different
- My key take aways:
 - Setting likely matters
 - Colonoscopy is effective...if completed
 - · The benefits of colonoscopy likely take time

Colonoscopy Prevalence (2000 to 2018) and CRC Incidence (2000 to 2016)



Helsingen, et al. NEJM Evid. 2022, 1. Bretthauer et al, N Engl J Med 2022; 387:1547-1556. Siegel et al, CA Cancer J Clin. 2020 May;70(3):145-164.

CRC Screening Should Begin at Age 45 (USPSTF Recommendations for Average-risk individuals)

U.S. Preventive Services TASK FORCE Colorectal Cancer: Screening Published Final Recommendations		Adults aged 50 to 75 years	The USPSTF recommends screening for colorectal cancer in all adults aged 50 to 75 years.	А
		Adults aged 45 to 49 years	The USPSTF recommends screening for colorectal cancer in adults aged 45 to 49 years.	В
		Adults aged 76 to 85 years	The USPSTF recommends that clinicians selectively offer screening for	
			colorectal cancer in adults aged 76 to 85 years. Evidence indicates that the net benefit of screening all persons in this age group is small. In determining whether this service is appropriate in individual cases,	С
	May 2021		patients and clinicians should consider the patient's overall health, prior screening history, and preferences.	

1 in 3 Americans are unscreened.1 in 2 are unscreened in under-resourced settings.



Davidson, K et al. JAMA. 2021; 325(19):1965-1977.

Age to Start Screening

Risk Group	Age to Start Screening	Age to Stop Screening	
Family history of CRC	40 <u>OR</u> 10 years before age of youngest family member diagnosed	Varies	
Inflammatory bowel disease; Familial polyposis syndrome; Hereditary colon cancer	Screen early (age varies)	Varies	
African-American/Black	45	Grade A: 75 Grade C: 85	
Average Risk	45	Grade A: 75 Grade C: 85	



Davidson, K et al. JAMA. 2021; 325(19):1965-1977.

2021 USPSTF CRC Recommendations

- Recommendations based on two commissioned reports:
 - 1) Systematic review of benefits and harms of screening adults 40 years or older:
 - a) Effectiveness and accuracy of screening tests
 - b) Comparative effectiveness of screening tests
 - c) Serious harms of different screening tests.
 - 2) Comparative modeling report from the CISNET Colorectal Cancer Working Group:
 - a) Life-years gained
 - b) CRC cases/deaths averted
 - c) Colonoscopy burden
 - d) Harms





USPSTF Recommended Screening Modalities (Average-risk individuals)



Direct-visualization techniques





Non-colonoscopic Screening Tests are Two-Step Strategies





Test Characteristics of Screening Tests

Test	Sensitivit y for CRC	Sensitivity for adv. Adenoma	Specificity for CRC	Evidenc e	Risk	Deaths averted per 1000 screened
High sensitivity guaiac FOBT	62- 79%	7%	87%-96%	Strong	Low	26
FIT	76- 95%	27%-47%	89%-96%	Weak	Low	26
FIT-DNA (Cologuard)	-DNA oguard) 93% 43%		85%	Early	Low	28 (yearly) 25 (Q 3 years)
CT Colonography	96%	67%-94% (>10mm) 73%-98% (6mm)	86%-98% (>10mm) 80%-93% (>6mm)	Weak	Low	26
Flexible58-Sigmoidoscopy75%		92%	Strong	Interm ediate	24 (28 with FIT)	
89%-98%			Knudsen e	t al. JAMA. 2	021; 325(19): :1998-201	

Patient Considerations for Screening Options

	HSgFOBT	FIT	FIT-DNA	CT Colonography	FS (+ FIT)	Colonoscopy
Invasiveness	+	+	+	++	++	+++
Home test	Yes	Yes	Yes	No	No	No
Dietary restrictions	Yes	No	No	Yes	Yes	Yes
Interval	1 year	1 year	1-3 years	5 years	5 (10 years)	10 years (if normal)
Complication s	Negligible	Negligible	Negligible	Few	Few	Low (0.1%)
Patient Participation	Moderate	Moderate	Moderate	Moderate	Moderate	Lowest
Cost	\$	\$	\$\$	\$\$	\$	\$\$



Robertson et al. AJG; 2017: 112; 37-53. Inadomi. NEJM; 2017; 376:1598-1600.

Emerging Non-Invasive Strategies (Not yet recommended for first-line average-risk)









New Stool-Based Tests Blood-Based/ MCED Wireless Capsule Colonoscopy Urine-based



High Risk Groups

- Personal history of tubular adenomas or CRC
- Family History of CRC/polyps
- Ulcerative colitis
- Crohn's disease





Screening Utilization and Barriers to Screening in the Underserved



CRC Outcomes Vary by Race/Ethnicity

Colorectal cancer incidence (2015–2019) by sex, race, and ethnicity; US

Colorectal cancer mortality (2016–2020) by sex, race, and ethnicity; US





1) Siegel RL, et al. CA Cancer J Clin. 2023; 2) North American Association of Central Cancer Registries, 2022. 3) National Center for Health Statistics, 2022.

CRC Outcomes Vary by Race/Ethnicity

CRC stage distribution in US by race/ethnicity; 2015-2019





Siegel RL, et al. CA Cancer J Clin. 2023.

CRC Outcomes Vary by Race/Ethnicity

CRC 5-year survival in US by race/ethnicity; 2012-2018





Siegel RL, et al. CA Cancer J Clin. 2023.

Screening Participation Contributes to Outcome Disparities by Race/Ethnicity





National Health Interview Survey, 2021.

Screening & Social Determinants of Health





National Health Interview Survey, 2021.

Screening & Social Determinants of Health





National Health Interview Survey, 2021.

Screening Rates are Low in Safety-Net Care Settings

U.S. adults age 50–75 years up-to-date with CRC screening



U.S. FQHC patients age 50-75 years up-to-date with CRC Screening





Centers for Disease Control and Prevention. Behavioral Risk Factor Surveillance System. 2012–2020. Uniform Data System: https://bphc.hrsa.gov/datareporting/reporting/index.html.

Social Determinants of Health





Barriers to Screening in the Medically Underserved

Patient-Level Factors

Lack of Knowledge Beliefs/Cultural factors Health Literacy Language Fear of procedure/prep Fear of cancer diagnosis Distrust Concerns re: provider quality Distance to endoscopy Cost/Lack of Insurance Comorbidities Competing demands Logistical challenges Lack of escort Time off work



Policy-Level Factors

Screening guidelines Insurance access Insurance mandate policy Coverage policy Cost/Co-pay policy

Williams R et al, Clin Transl Gastroent. 2016. White P, Itzkowitz S. Curr Gastro Rep, 2020. Carethers JM, Doubeni CA. Gastro, 2020. May FP et al, J Ca Educ, 2016. May FP et al. Am J Gastroenterol, 2015. May FP et al. Med Care, 2019.

Using Implementation Science to Increase Screening Participation and Eliminate Disparities





Taplin SH et al. J Natl Cancer Inst Monogr. 2012;2012(44):2-10. Carethers JM, Doubeni CA. Gastro. 2020 Jan;158(2):354-367.

Patient-focused Intervention: Navigation

Patients, setting: Low-income Black and Latino individuals age 50-75 years. One large medical center in Boston. N=843

Design: RCT.

Intervention

Arm 1: Telephone-delivered individualized education by two bilingual navigators. Arm 2: Usual care

Outcome: Colonoscopy completion within 6 months





DeGroff A, et al. Am J Prev Med. 2017 Sep;53(3):363-372.

System-level Intervention: Mailed FIT



Patients, setting: Safety-net system (8 clinics); Majority Black and Latino patients age 50-75 years. N=10,820.

Design: Cluster randomized trial

Intervention

Arm 1: Mailed postcard + telephone call + mailed FIT kit + Reminder call Arm 2: Usual care

Outcome: Screening participation at 1 year



Somsouk M et al. J Natl Cancer Inst. 2020 Mar 1;112(3):305-313.

System-level: Screening improvements reduce incidence and mortality disparities

Patients, setting: Kaiser Permanente Northern California health plan patients; 11% Black individuals; age 50-75 years. N=792,081.

Design: Retrospective cohort

Intervention

Organized, multilevel screening program fully implemented by 2008. Patients followed through age 79.

Outcome: Screening participation at 1 year

CRC incidence & mortality by race, 2000–2019





Doubeni CA et al., N Engl J Med 2022; 386:796-798.

Beyond Screening Interventions





Policy • Community Engagement • Dissemination Throughout the cancer care continuum

Diet Modification

- Minimize processed meats:
 - Ham, bacon, hot dogs, raw sausages (salami), bologna, blood sausage, pate, meat spreads, cold cuts, canned meats, corned beef
- Minimize red meats:
 - Beef, pork, lamb, goat
- Increase intake of:
 - Whole grains, fiber, fruit, non-starchy vegetables, vitamin C-rich foods, fish, vitamin D







Slide credit: UCLA Digestive Health Nutrition Program

Other Lifestyle Changes

- Obesity and diabetes prevention
- Increase physical activity
- Drink alcohol in moderation
 - Maximum of 2 drinks/day for men; 1 drink/day for women
 - Maximum 14 units/week for men and women
- Avoid tobacco





Key Take Away Points

- CRC is common and deadly but is largely preventable with screening.
- Screening for CRC is evidence-based and recommended for all adults but underutilized.
- Screening should begin at 45 for average-risk individuals (& earlier in high-risk groups).
- There are several recommended screening modalities, and patients should select the modality most appropriate for them.
- Additional prevention strategies include diet modification and avoidance of obesity, tobacco, and heavy alcohol.
- Disparities in screening utilization contribute to inequities in CRC outcomes and warrant tailored, targeted interventions.





Frequently Asked Questions (FAQs)

1. What is the USPSTF recommended age to initiate colorectal cancer screening in average-risk adults?

Age 45. This change was made in May of 2021.

2. When should individuals with a family history of colorectal cancer initiate screening?

40 OR 10 years before age of youngest family member diagnosed

3. What is the best colorectal cancer screening test for average-risk individuals?

Any of the 7 recommended by USPSTF. "The best test is the test that gets done"

4. What is the appropriate age to stop colorectal cancer screening? Grade A: 75; Grade C: 85



Thank You!



Vatche and Tamar Manoukian Division of Digestive Diseases

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U.S. Department of Veterans Affairs

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#MayLabUCLA

https:/www.uclahealth.org/gastro/may-lab

X@drfolamay



Q & A Session



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Thank you!